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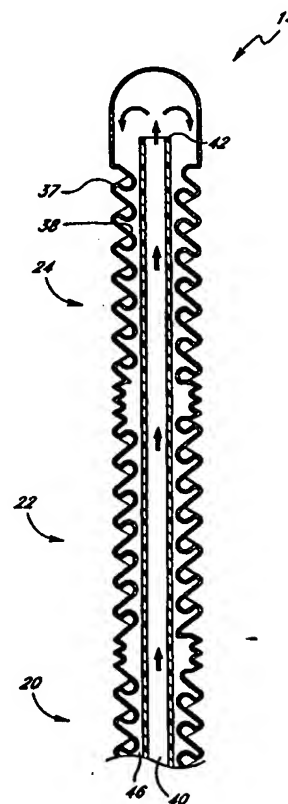
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(54) Title: SELECTIVE ORGAN COOLING APPARATUS AND METHOD

(57) Abstract

This invention is a selective organ heat transfer device with deep irregularities in a turbulence inducing exterior surface. The device can have a plurality of elongated, articulated segments (20) each having a turbulence inducing exterior surface. A flexible joint (25) connects adjacent elongated, articulated segments. An inner lumen is disposed within the heat transfer segments. The inner lumen (42) is capable of transporting a pressurized working fluid to a distal end of the heat transfer element.



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## TITLE OF THE INVENTION

Selective Organ Cooling Apparatus  
and Method

## 5 BACKGROUND OF THE INVENTION

Field of the Invention - The present invention relates generally to the modification and control of the temperature of a selected body organ. More particularly, the invention relates to a method and intravascular apparatus for controlling organ temperature.

10 Background Information - Organs in the human body, such as the brain, kidney and heart, are maintained at a constant temperature of approximately 37° C. Hypothermia can be clinically defined as a core body temperature of 35° C or less. Hypothermia is sometimes characterized further according to its severity. A body core temperature in the range of 33° C to 35° C is described as mild hypothermia. A body  
15 temperature of 28° C to 32° C is described as moderate hypothermia. A body core temperature in the range of 24° C to 28° C is described as severe hypothermia.

Hypothermia is uniquely effective in reducing brain injury caused by a variety of neurological insults and may eventually play an important role in emergency brain resuscitation. Experimental evidence has demonstrated that cerebral cooling improves  
20 outcome after global ischemia, focal ischemia, or traumatic brain injury. For this reason, hypothermia may be induced in order to reduce the effect of certain bodily injuries to the brain as well as other organs.

Cerebral hypothermia has traditionally been accomplished through whole body cooling to create a condition of total body hypothermia in the range of 20° C to 30° C.  
25 However, the use of total body hypothermia risks certain deleterious systematic vascular effects. For example, total body hypothermia may cause severe derangement of the cardiovascular system, including low cardiac output, elevated systematic resistance, and ventricular fibrillation. Other side effects include renal failure, disseminated intravascular coagulation, and electrolyte disturbances. In addition to the undesirable  
30 side effects, total body hypothermia is difficult to administer.

Catheters have been developed which are inserted into the bloodstream of the patient in order to induce total body hypothermia. For example, U.S. Patent No.

3,425,419 to Dato describes a method and apparatus of lowering and raising the temperature of the human body. Dato induces moderate hypothermia in a patient using a metallic catheter. The metallic catheter has an inner passageway through which a fluid, such as water, can be circulated. The catheter is inserted through the femoral vein and then through the inferior vena cava as far as the right atrium and the superior vena cava. The Dato catheter has an elongated cylindrical shape and is constructed from stainless steel. By way of example, Dato suggests the use of a catheter approximately 70 cm in length and approximately 6 mm in diameter. However, use of the Dato device implicates the negative effects of total body hypothermia described above.

Due to the problems associated with total body hypothermia, attempts have been made to provide more selective cooling. For example, cooling helmets or head gear have been used in an attempt to cool only the head rather than the patient's entire body. However, such methods rely on conductive heat transfer through the skull and into the brain. One drawback of using conductive heat transfer is that the process of reducing the temperature of the brain is prolonged. Also, it is difficult to precisely control the temperature of the brain when using conduction due to the temperature gradient that must be established externally in order to sufficiently lower the internal temperature. In addition, when using conduction to cool the brain, the face of the patient is also subjected to severe hypothermia, increasing discomfort and the likelihood of negative side effects. It is known that profound cooling of the face can cause similar cardiovascular side effects as total body cooling. From a practical standpoint, such devices are cumbersome and may make continued treatment of the patient difficult or impossible.

Selected organ hypothermia has been accomplished using extracorporeal perfusion, as detailed by Arthur E. Schwartz, M.D. et al., in *Isolated Cerebral Hypothermia by Single Carotid Artery Perfusion of Extracorporeally Cooled Blood in Baboons*, which appeared in Vol. 39, No. 3, NEUROSURGERY 577 (September, 1996). In this study, blood was continually withdrawn from baboons through the femoral artery. The blood was cooled by a water bath and then infused through a common carotid artery with its external branches occluded. Using this method, normal heart rhythm, systemic arterial blood pressure and arterial blood gas values were maintained during the hypothermia. This study showed that the brain could be selectively cooled to temperatures of 20° C without reducing the temperature of the entire body. However,

external circulation of blood is not a practical approach for treating humans because the risk of infection, need for anticoagulation, and risk of bleeding is too great. Further, this method requires cannulation of two vessels making it more cumbersome to perform particularly in emergency settings. Even more, percutaneous cannulation of the carotid artery is difficult and potentially fatal due to the associated arterial wall trauma. Finally, this method would be ineffective to cool other organs, such as the kidneys, because the feeding arteries cannot be directly cannulated percutaneously.

Selective organ hypothermia has also been attempted by perfusion of a cold solution such as saline or perfluorocarbons. This process is commonly used to protect the heart during heart surgery and is referred to as cardioplegia. Perfusion of a cold solution has a number of drawbacks, including a limited time of administration due to excessive volume accumulation, cost, and inconvenience of maintaining the perfusate and lack of effectiveness due to the temperature dilution from the blood. Temperature dilution by the blood is a particular problem in high blood flow organs such as the brain.

Therefore, a practical method and apparatus which modifies and controls the temperature of a selected organ satisfies a long-felt need.

#### BRIEF SUMMARY OF THE INVENTION

The apparatus of the present invention can, by way of example only, include a heat transfer element which comprises first and second elongated, articulated segments, each segment having a turbulence-inducing exterior surface. A flexible joint can connect the first and second elongated segments. An inner coaxial lumen may be disposed within the first and second elongated segments and is capable of transporting a pressurized working fluid to a distal end of the first elongated segment. In addition, the first and second elongated segments may have a turbulence-inducing interior surface for inducing turbulence within the pressurized working fluid. The turbulence-inducing exterior surface may be adapted to induce turbulence within a free stream of blood flow when placed within an artery. The turbulence-inducing exterior surface may be adapted to induce a turbulence intensity greater than 0.05 within a free stream blood flow. In one embodiment, the flexible joint comprises a bellows section which also allows for axial compression of the heat transfer element.

In one embodiment, the turbulence-inducing exterior surfaces of the heat transfer element comprise one or more helical ridges configured to have a depth which is greater than a thickness of a boundary layer of blood which develops within an arterial blood flow. Adjacent segments of the heat transfer element can be oppositely spiraled to increase turbulence. For instance, the first elongated heat transfer segment may comprise one or more helical ridges having a counter-clockwise twist, while the second elongated heat transfer segment comprises one or more helical ridges having a clockwise twist. Alternatively, of course, the first elongated heat transfer segment may comprise one or more clockwise helical ridges, and the second elongated heat transfer segment may comprise one or more counter-clockwise helical ridges. The first and second elongated, articulated segments may be formed from highly conductive materials.

In another embodiment, the turbulence-inducing exterior surface of the heat transfer element is adapted to induce turbulence throughout the duration of each pulse of a pulsatile blood flow when placed within an artery. In still another embodiment, the turbulence-inducing exterior surface of the heat transfer element is adapted to induce turbulence during at least 20% of the period of each cardiac cycle when placed within an artery.

The heat transfer device may also have a coaxial supply catheter with an inner catheter lumen coupled to the inner coaxial lumen within the first and second elongated heat transfer segments. A working fluid supply configured to dispense the pressurized working fluid may be coupled to the inner catheter lumen. The working fluid supply may be configured to produce the pressurized working fluid at a temperature of about 0°C and at a pressure below about 5 atmospheres of pressure.

In yet another alternative embodiment, the heat transfer device may have three or more elongated, articulated, heat transfer segments having a turbulence-inducing exterior surface, with additional flexible joints connecting the additional elongated heat transfer segments. In one such embodiment, by way of example, the first and third elongated heat transfer segments may comprise clockwise helical ridges, and the second elongated heat transfer segment may comprise one or more counter-clockwise helical ridges. Alternatively, of course, the first and third elongated heat transfer segments may comprise counter-clockwise helical ridges, and the second elongated heat transfer-segment may comprise one or more clockwise helical ridges.

The turbulence-inducing exterior surface of the heat transfer element may optionally include a surface coating or treatment to inhibit clot formation. One variation of the heat transfer element comprises a stent coupled to a distal end of the first elongated heat transfer segment.

5       The present invention also envisions a method of treating the brain which comprises the steps of inserting a flexible, conductive heat transfer element into a carotid artery from a distal location, and circulating a working fluid through the flexible, conductive heat transfer element in order to selectively modify the temperature of the brain without significantly modifying the temperature of the entire body. The flexible,  
10       conductive heat transfer element preferably absorbs more than about 25, 50 or 75 Watts of heat.

      The method may also comprise the step of inducing turbulence within the free stream blood flow within the carotid artery. In one embodiment, the method includes the step of inducing blood turbulence with a turbulence intensity greater than about 0.05  
15       within the carotid artery. In another embodiment, the method includes the step of inducing blood turbulence throughout the duration of the period of the cardiac cycle within the carotid artery. In yet another embodiment, the method comprises the step of inducing blood turbulence throughout the period of the cardiac cycle within the carotid artery or during greater than about 20% of the period of the cardiac cycle within the  
20       carotid artery. The step of circulating may comprise the step of inducing turbulent flow of the working fluid through the flexible, conductive heat transfer element. The pressure of the working fluid may be maintained below about 5 atmospheres of pressure.

      The present invention also envisions a method for selectively cooling an organ in the body of a patient which comprises the steps of introducing a catheter, with a heat  
25       transfer element, into a blood vessel supplying the organ, the catheter having a diameter of about 4 mm or less, inducing free stream turbulence in blood flowing over the heat transfer element, and cooling the heat transfer element to remove heat from the blood to cool the organ without substantially cooling the entire body. In one embodiment, the cooling step removes at least about 75 Watts of heat from the blood. In another  
30       embodiment, the cooling step removes at least about 100 Watts of heat from the blood. The organ being cooled may be the human brain.

The step of inducing free stream turbulence may induce a turbulence intensity greater than about 0.05 within the blood vessel. The step of inducing free stream turbulence may induce turbulence throughout the duration of each pulse of blood flow. The step of inducing free stream turbulence may induce turbulence for at least about 20%  
5 of the duration of each pulse of blood flow.

In one embodiment, the catheter has a flexible metal tip and the cooling step occurs at the tip. The tip may have turbulence-inducing elongated heat transfer segments separated by bellows sections. The turbulence-inducing segments may comprise helical ridges which are configured to have a depth which is greater than a thickness of a  
10 boundary layer of blood which develops within the blood vessel. In another embodiment, the catheter has a tip at which the cooling step occurs and the tip has turbulence-inducing elongated heat transfer segments that alternately spiral bias the surrounding blood flow in clockwise and counterclockwise directions.

The cooling step may comprise the step of circulating a working fluid in through  
15 an inner lumen in the catheter and out through an outer, coaxial lumen. In one embodiment, the working fluid remains a liquid throughout the cycle. The working fluid may be aqueous.

The present invention also envisions a cooling catheter comprising a catheter shaft having first and second lumens therein. The cooling catheter also comprises a cooling tip  
20 adapted to transfer heat to or from a working fluid circulated in through the first lumen and out through the second lumen, and turbulence-inducing structures on the cooling tip capable of inducing free stream turbulence when the tip is inserted into a blood vessel. The turbulence-inducing structures may induce a turbulence intensity of at least about 0.05. The cooling tip may be adapted to induce turbulence within the working fluid. The  
25 catheter is capable of removing at least about 25 Watts of heat from an organ when inserted into a vessel supplying that organ, while cooling the tip with a working fluid that remains a liquid in the catheter. Alternatively, the catheter is capable of removing at least about 50 or 75 Watts of heat from an organ when inserted into a vessel supplying that organ, while cooling the tip with an aqueous working fluid. In one embodiment, in  
30 use, the tip has a diameter of about 4 mm or less. Optionally, the turbulence-inducing surfaces on the heat transfer segments comprise helical ridges which have a depth sufficient to disrupt the free stream blood flow in the blood vessel. Alternatively, the



turbulence-inducing surfaces may comprise staggered protrusions from the outer surfaces of the heat transfer segments, which have a height sufficient to disrupt the free stream flow of blood within the blood vessel.

In another embodiment, a cooling catheter may comprise a catheter shaft having  
5 first and second lumens therein, a cooling tip adapted to transfer heat to or from a  
working fluid circulated in through the first lumen and out through the second lumen,  
and turbulence-inducing structures on the cooling tip capable of inducing turbulence  
when the tip is inserted into a blood vessel. Alternatively, a cooling catheter may  
comprise a catheter shaft having first and second lumens therein, a cooling tip adapted to  
10 transfer heat to or from a working fluid circulated in through the first lumen and out  
through the second lumen, and structures on the cooling tip capable of inducing free  
stream turbulence when the tip is inserted into a blood vessel. In another embodiment, a  
cooling catheter may comprise a catheter shaft having first and second lumens therein, a  
cooling tip adapted to transfer heat to or from a working fluid circulated in through the  
15 first lumen and out through the second lumen, and turbulence-inducing structures on the  
cooling tip capable of inducing turbulence with an intensity greater than about 0.05 when  
the tip is inserted into a blood vessel.

The novel features of this invention, as well as the invention itself, will be best  
20 understood from the attached drawings, taken along with the following description, in  
which similar reference characters refer to similar parts, and in which:

#### BRIEF DESCRIPTION OF THE SEVERAL VIEWS OF THE DRAWINGS

Figure 1 is a graph illustrating the velocity of steady state turbulent flow as a  
25 function of time;

Figure 2A is a graph showing the velocity of the blood flow within an artery as a  
function of time;

Figure 2B is a graph illustrating the velocity of steady state turbulent flow under  
pulsatile conditions as a function of time, similar to arterial blood flow;

30 Figure 2C is an elevation view of a turbulence inducing heat transfer element  
within an artery;

Figure 3A is a velocity profile diagram showing a typical steady state Poiseuillean flow driven by a constant pressure gradient;

Figure 3B is a velocity profile diagram showing blood flow velocity within an artery, averaged over the duration of the cardiac pulse;

Figure 3C is a velocity profile diagram showing blood flow velocity within an artery, averaged over the duration of the cardiac pulse, after insertion of a smooth heat transfer element within the artery;

Figure 4 is an elevation view of one embodiment of a heat transfer element according to the invention;

10 Figure 5 is longitudinal section view of the heat transfer element of Figure 4;

Figure 6 is a transverse section view of the heat transfer element of Figure 4;

Figure 7 is a perspective view of the heat transfer element of Figure 4 in use within a blood vessel;

Figure 8 is a cut-away perspective view of an alternative embodiment of a heat  
15 transfer element according to the invention;

Figure 9 is a transverse section view of the heat transfer element of Figure 8; and

Figure 10 is a schematic representation of the invention being used in one embodiment to cool the brain of a patient.

20 DETAILED DESCRIPTION OF THE INVENTION

In order to intravascularly regulate the temperature of a selected organ, a heat transfer element may be placed in the feeding artery of the organ to absorb or deliver the heat from or to the blood flowing into the organ. The transfer of heat may cause either a cooling or a heating of the selected organ. The heat transfer element must be small enough to fit within the feeding artery while still allowing a sufficient blood flow to reach the organ in order to avoid ischemic organ damage. A heat transfer element which selectively cools an organ should be capable of providing the necessary heat transfer rate to produce the desired cooling or heating effect within the organ. By placing the heat transfer element within the feeding artery of an organ, the temperature of an organ can be controlled without significantly affecting the remaining parts of the body. These points can be illustrated by using brain cooling as an example.

The common carotid artery supplies blood to the head and brain. The internal carotid artery branches off of the common carotid to directly supply blood to the brain. To selectively cool the brain, the heat transfer element is placed into the common carotid artery, or both the common carotid artery and the internal carotid artery. The internal  
5 diameter of the common carotid artery ranges from 6 to 8 mm and the length ranges from 80 to 120 mm. Thus, the heat transfer element residing in one of these arteries cannot be much larger than 4 mm in diameter in order to avoid occluding the vessel.

It is important that the heat transfer element be flexible in order to be placed within the small feeding artery of an organ. Feeding arteries, like the carotid artery,  
10 branch off the aorta at various levels. Subsidiary arteries continue to branch off the initial branches. For example, the internal carotid artery is a small diameter artery that branches off of the common carotid artery near the angle of the jaw. Because the heat transfer element is typically inserted into a peripheral artery, such as the femoral artery, and accesses the feeding artery by initially passing through a series of one or more of  
15 these branches, the flexibility of the heat transfer element is an important characteristic of the heat transfer element. Further, the heat transfer element is ideally constructed from a highly thermally conductive material such as metal in order to facilitate heat transfer. The use of a highly thermally conductive material increases the heat transfer rate for a given temperature differential between the coolant within the heat transfer element and  
20 the blood. This facilitates the use of a higher temperature coolant within the heat transfer element, allowing safer coolants, such as water, to be used. Highly thermally conductive materials, such as metals, tend to be rigid. Therefore, the design of the heat transfer element should facilitate flexibility in an inherently inflexible material.

In order to obtain the benefits of hypothermia described above, it is desirable to  
25 reduce the temperature of the blood flowing to the brain to between 30° C and 32° C. Given that a typical brain has a blood flow rate through each carotid artery (right and left) of approximately 250-375 cubic centimeters per minute, the heat transfer element should absorb 75-175 Watts of heat when placed in one of the carotid arteries, in order to induce the desired cooling effect. It should be noted that smaller organs may have less blood  
30 flow in the supply artery and may require less heat transfer, such as 25 Watts.

When a heat transfer element is inserted coaxially into an artery, the primary mechanism of heat transfer between the surface of the heat transfer element and the

blood is forced convection. Convection relies upon the movement of fluid to transfer heat. Forced convection results when an external force causes motion within the fluid. In the case of arterial flow, the beating heart causes the motion of the blood around the heat transfer element.

5       The magnitude of the heat transfer rate is proportional to the surface area of the heat transfer element, the temperature differential, and the heat transfer coefficient of the heat transfer element.

As noted above, the receiving artery into which the heat transfer element is placed has a limited diameter and length. Thus, surface area of the heat transfer element must  
10       be limited, to avoid significant obstruction of the artery, and to allow the heat transfer element to easily pass through the vascular system. For placement within the internal and common carotid artery, the cross sectional diameter of the heat transfer element is limited to about 4 mm, and its length is limited to approximately 10 cm.

The temperature differential can be increased by decreasing the surface  
15       temperature of the heat transfer element. However, the minimum allowable surface temperature is limited by the characteristics of blood. Blood freezes at approximately 0° C. When the blood approaches freezing, ice emboli may form in the blood which may lodge downstream, causing serious ischemic injury. Furthermore, reducing the temperature of the blood also increases its viscosity, which results in a small decrease in  
20       the value of the convection heat transfer coefficient. In addition, increased viscosity of the blood may result in an increase in the pressure drop within the artery, thus, compromising the flow of blood to the brain. Given the above constraints, it is advantageous to limit the minimum allowable surface temperature of the heat transfer element to approximately 5° C. This results in a maximum temperature differential  
25       between the blood stream and the heat transfer element of approximately 32° C.

The mechanisms by which the value of the convection heat transfer coefficient may be increased are complex. However, it is well known that the convection heat transfer coefficient increases with the level of turbulent kinetic energy in the fluid flow. Thus it is advantageous to have turbulent blood flow in contact with the heat transfer  
30       element.

Figure 1 is a graph illustrating steady state turbulent flow. The vertical axis is the velocity of the flow. The horizontal axis represents time. The average velocity of the

turbulent flow is shown by a line 100. The actual instantaneous velocity of the flow is shown by a curve 102.

Under constant pressure conditions, steady flows in pipes are characterized as a balance between viscous stresses and the constant pressure gradient. Such flows are called Poiseuillean. Figure 3A is a velocity profile diagram showing a typical steady state Poiseuillean flow driven by a constant pressure gradient. The velocity of the fluid across the pipe is shown in Figure 3A by the parabolic curve and corresponding velocity vectors. The velocity of the fluid in contact with the wall of the pipe is zero. The boundary layer is the region of the flow in contact with the pipe surface in which viscous stresses are dominant. In steady state Poiseuillean flow, the boundary layer develops until it includes the whole pipe, i.e., the boundary layer thickness in Figure 3A is one half of the diameter of the pipe.

Under conditions of Poiseuillean flow, the Reynolds number, the ratio of inertial forces to viscous forces, can be used to characterize the level of turbulent kinetic energy existing in the flow. For Poiseuillean flows, Reynolds numbers must be greater than about 2300 to cause a transition from laminar to turbulent flow. Further, when the Reynolds number is greater than about 2000, the boundary layer is receptive to "tripping". Tripping is a process by which a small perturbation in the boundary layer can create turbulent conditions. The receptivity of a boundary layer to "tripping" is proportional to the Reynolds number and is nearly zero for Reynolds numbers less than 2000.

In contrast with the steady Poiseuillean flow, the blood flow in arteries is induced by the beating heart and is therefore pulsatile. Figure 2A is a graph showing the velocity of the blood flow within an artery as a function of time. The beating heart provides pulsatile flow with an approximate period of 0.5 to 1 second. This is known as the period of the cardiac cycle. The horizontal axis in Figure 2A represents time in seconds and the vertical axis represents the average velocity of blood in centimeters per second. Although very high velocities are reached at the peak of the pulse, the high velocity occurs for only a small portion of the cycle. In fact, the velocity of the blood reaches zero in the carotid artery at the end of a pulse and temporarily reverses.

Because of the relatively short duration of the cardiac pulse, the blood flow in the arteries does not develop into the classic Poiseuillean flow. Figure 3B is a velocity

profile diagram showing blood flow velocity within an artery averaged over the cardiac pulse. The majority of the flow within the artery has the same velocity. The boundary layer where the flow velocity decays from the free stream value to zero is very thin, typically  $1/6$  to  $1/20$  of the diameter of the artery, as opposed to one half of the diameter of the artery in the Poiseuillean flow condition.

As noted above, if the flow in the artery were steady rather than pulsatile, the transition from laminar to turbulent flow would occur when the value of the Reynolds number exceeds about 2,000. However, in the pulsatile arterial flow, the value of the Reynolds number varies during the cardiac cycle, just as the flow velocity varies. In pulsatile flows, due to the enhanced stability associated with the acceleration of the free stream flow, the critical value of the Reynolds number at which the unstable modes of motion grow into turbulence is found to be much higher, perhaps as high as 9,000.

The blood flow in the arteries of interest remains laminar over more than 80% of the cardiac cycle. Referring again to Figure 2A, the blood flow is turbulent from approximately time  $t_1$  until time  $t_2$  during a small portion of the descending systolic flow, which is less than 20% of the period of the cardiac cycle. If a heat transfer element is placed inside the artery, heat transfer will be facilitated during this short interval. However, to transfer the necessary heat to cool the brain, turbulent kinetic energy should be produced in the blood stream and sustained throughout the entire period of the cardiac cycle.

A thin boundary layer has been shown to form during the cardiac cycle. This boundary layer will form over the surface of a smooth heat transfer element. Figure 3C is a velocity profile diagram showing blood flow velocity within an artery, averaged over the cardiac pulse, after insertion of a smooth heat transfer element within the artery. In Figure 3C, the diameter of the heat transfer element is about one half of the diameter of the artery. Boundary layers develop adjacent to the heat transfer element as well as next to the walls of the artery. Each of these boundary layers has approximately the same thickness as the boundary layer which would have developed at the wall of the artery in the absence of the heat transfer element. The free stream flow region is developed in an annular ring around the heat transfer element.

One way to increase the heat transfer rate is to create a turbulent boundary layer on the heat transfer element surface. However, turbulence in the very thin boundary layer

will not produce sufficient kinetic energy to produce the necessary heat transfer rate. Therefore, to induce sufficient turbulent kinetic energy to increase the heat transfer rate sufficiently to cool the brain, a stirring mechanism, which abruptly changes the direction of velocity vectors, should be utilized. This can create high levels of turbulence intensity in the free stream, thereby sufficiently increasing the heat transfer rate.

This turbulence intensity should ideally be sustained for a significant portion of the cardiac cycle. Further, turbulent kinetic energy should ideally be created throughout the free stream and not just in the boundary layer. Figure 2B is a graph illustrating the velocity of continually turbulent flow under pulsatile conditions as a function of time, which would result in optimal heat transfer in arterial blood flow. Turbulent velocity fluctuations are seen throughout the cycle as opposed to the short interval of fluctuations seen in Figure 2A between time  $t_1$  and time  $t_2$ . These velocity fluctuations are found within the free stream. The turbulence intensity shown in Figure 2B is at least 0.05. In other words, the instantaneous velocity fluctuations deviate from the mean velocity by at least 5%. Although, ideally, turbulence is created throughout the entire period of the cardiac cycle, the benefits of turbulence are also obtained if the turbulence is sustained for only 75%, 50% or even as low as 30% or 20% of the cardiac cycle.

To create the desired level of turbulence intensity in the blood free stream during the whole cardiac cycle, one embodiment of the invention uses a modular design. This design creates helical blood flow and produces a high level of turbulence in the free stream by periodically forcing abrupt changes in the direction of the helical blood flow. Figure 2C is a perspective view of such a turbulence inducing heat transfer element within an artery. Turbulent flow would be found at point 114, in the free stream area. The abrupt changes in flow direction are achieved through the use of a series of two or more heat transfer segments, each comprised of one or more helical ridges. To affect the free stream, the depth of the helical ridge is larger than the thickness of the boundary layer which would develop if the heat transfer element had a smooth cylindrical surface.

The use of periodic abrupt changes in the helical direction of the blood flow in order to induce strong free stream turbulence may be illustrated with reference to a common clothes washing machine. The rotor of a washing machine spins initially in one direction causing laminar flow. When the rotor abruptly reverses direction, significant

turbulent kinetic energy is created within the entire wash basin as the changing currents cause random turbulent motion within the clothes-water slurry.

Figure 4 is an elevation view of one embodiment of a heat transfer element 14 according to the present invention. The heat transfer element 14 is comprised of a series of elongated, articulated segments or modules 20, 22, 24. Three such segments are shown in this embodiment, but two or more such segments could be used without departing from the spirit of the invention. As seen in Figure 4, a first elongated heat transfer segment 20 is located at the proximal end of the heat transfer element 14. A turbulence-inducing exterior surface of the segment 20 comprises four parallel helical ridges 28 with four parallel helical grooves 26 therebetween. One, two, three, or more parallel helical ridges 28 could also be used without departing from the spirit of the present invention. In this embodiment, the helical ridges 28 and the helical grooves 26 of the heat transfer segment 20 have a left hand twist, referred to herein as a counter-clockwise spiral or helical rotation, as they proceed toward the distal end of the heat transfer segment 20.

The first heat transfer segment 20 is coupled to a second elongated heat transfer segment 22 by a first bellows section 25, which provides flexibility and compressibility. The second heat transfer segment 22 comprises one or more helical ridges 32 with one or more helical grooves 30 therebetween. The ridges 32 and grooves 30 have a right hand, or clockwise, twist as they proceed toward the distal end of the heat transfer segment 22. The second heat transfer segment 22 is coupled to a third elongated heat transfer segment 24 by a second bellows section 27. The third heat transfer segment 24 comprises one or more helical ridges 36 with one or more helical grooves 34 therebetween. The helical ridge 36 and the helical groove 34 have a left hand, or counter-clockwise, twist as they proceed toward the distal end of the heat transfer segment 24. Thus, successive heat transfer segments 20, 22, 24 of the heat transfer element 14 alternate between having clockwise and counterclockwise helical twists. The actual left or right hand twist of any particular segment is immaterial, as long as adjacent segments have opposite helical twist.

In addition, the rounded contours of the ridges 28, 32, 36 also allow the heat transfer element 14 to maintain a relatively atraumatic profile, thereby minimizing the



possibility of damage to the blood vessel wall. A heat transfer element according to the present invention may be comprised of two, three, or more heat transfer segments.

5 The bellows sections 25, 27 are formed from seamless and nonporous materials, such as metal, and therefore are impermeable to gas, which can be particularly important, depending on the type of working fluid which is cycled through the heat transfer element 14. The structure of the bellows sections 25, 27 allows them to bend, extend and compress, which increases the flexibility of the heat transfer element 14 so that it is more readily able to navigate through blood vessels. The bellows sections 25, 27 also provide for axial compression of the heat transfer element 14, which can limit the trauma when  
10 the distal end of the heat transfer element 14 abuts a blood vessel wall. The bellows sections 25, 27 are also able to tolerate cryogenic temperatures without a loss of performance.

The exterior surfaces of the heat transfer element 14 can be made from metal, and may comprise very high thermal conductivity materials such as nickel, thereby  
15 facilitating heat transfer. Alternatively, other metals such as stainless steel, titanium, aluminum, silver, copper and the like, can be used, with or without an appropriate coating or treatment to enhance biocompatibility or inhibit clot formation. Suitable biocompatible coatings include, e.g., gold, platinum or polymer paralyene. The heat transfer element 14 may be manufactured by plating a thin layer of metal on a mandrel  
20 that has the appropriate pattern. In this way, the heat transfer element 14 may be manufactured inexpensively in large quantities, which is an important feature in a disposable medical device.

Because the heat transfer element 14 may dwell within the blood vessel for extended periods of time, such as 24-48 hours or even longer, it may be desirable to treat  
25 the surfaces of the heat transfer element 14 to avoid clot formation. In particular, one may wish to treat the bellows sections 25, 27 because stagnation of the blood flow may occur in the convolutions, thus allowing clots to form and cling to the surface to form a thrombus. One means by which to prevent thrombus formation is to bind an antithrombogenic agent to the surface of the heat transfer element 14. For example,  
30 heparin is known to inhibit clot formation and is also known to be useful as a biocoating. Alternatively, the surfaces of the heat transfer element 14 may be bombarded with ions

such as nitrogen. Bombardment with nitrogen can harden and smooth the surface and, thus prevent adherence of clotting factors to the surface.

Figure 5 is a longitudinal sectional view of the heat transfer element 14 of an embodiment of the invention, taken along line 5-5 in Figure 4. Some interior contours are omitted for purposes of clarity. An inner tube 42 creates an inner coaxial lumen 42 and an outer coaxial lumen 46 within the heat transfer element 14. Once the heat transfer element 14 is in place in the blood vessel, a working fluid such as saline or other aqueous solution may be circulated through the heat transfer element 14. Fluid flows up a supply catheter into the inner coaxial lumen 40. At the distal end of the heat transfer element 14, the working fluid exits the inner coaxial lumen 40 and enters the outer lumen 46. As the working fluid flows through the outer lumen 46, heat is transferred from the working fluid to the exterior surface 37 of the heat transfer element 14. Because the heat transfer element 14 is constructed from a high conductivity material, the temperature of its exterior surface 37 may reach very close to the temperature of the working fluid. The tube 42 may be formed as an insulating divider to thermally separate the inner lumen 40 from the outer lumen 46. For example, insulation may be achieved by creating longitudinal air channels in the wall of the insulating tube 42. Alternatively, the insulating tube 42 may be constructed of a non-thermally conductive material like polytetrafluoroethylene or some other polymer.

It is important to note that the same mechanisms that govern the heat transfer rate between the exterior surface 37 of the heat transfer element 14 and the blood also govern the heat transfer rate between the working fluid and the interior surface 38 of the heat transfer element 14. The heat transfer characteristics of the interior surface 38 are particularly important when using water, saline or other fluid which remains a liquid as the coolant. Other coolants such as freon undergo nucleate boiling and create turbulence through a different mechanism. Saline is a safe coolant because it is non-toxic, and leakage of saline does not result in a gas embolism, which could occur with the use of boiling refrigerants. Since turbulence in the coolant is enhanced by the shape of the interior surface 38 of the heat transfer element 14, the coolant can be delivered to the heat transfer element 14 at a warmer temperature and still achieve the necessary heat transfer rate.

This has a number of beneficial implications in the need for insulation along the catheter shaft length. Due to the decreased need for insulation, the catheter shaft diameter can be made smaller. The enhanced heat transfer characteristics of the interior surface of the heat transfer element 14 also allow the working fluid to be delivered to the heat transfer element 14 at lower flow rates and lower pressures. High pressures may make the heat transfer element stiff and cause it to push against the wall of the blood vessel, thereby shielding part of the exterior surface 37 of the heat transfer element 14 from the blood. Because of the increased heat transfer characteristics achieved by the alternating helical ridges 28, 32, 36, the pressure of the working fluid may be as low as 5 atmospheres, 3 atmospheres, 2 atmospheres or even less than 1 atmosphere.

Figure 6 is a transverse sectional view of the heat transfer element 14 of the invention, taken at a location denoted by the line 6-6 in Figure 4. Figure 6 illustrates a five lobed embodiment, whereas Figure 4 illustrates a four-lobed embodiment. As mentioned earlier, any number of lobes might be used. In Figure 6, the coaxial construction of the heat transfer element 14 is clearly shown. The inner coaxial lumen 40 is defined by the insulating coaxial tube 42. The outer lumen 46 is defined by the exterior surface of the insulating coaxial tube 42 and the interior surface 38 of the heat transfer element 14. In addition, the helical ridges 32 and helical grooves 30 may be seen in Figure 6. As noted above, in the preferred embodiment, the depth of the grooves,  $d_g$ , is greater than the boundary layer thickness which would have developed if a cylindrical heat transfer element were introduced. For example, in a heat transfer element 14 with a 4 mm outer diameter, the depth of the invaginations,  $d_g$ , may be approximately equal to 1 mm if designed for use in the carotid artery. Although Figure 6 shows four ridges and four grooves, the number of ridges and grooves may vary. Thus, heat transfer elements with 1, 2, 3, 4, 5, 6, 7, 8 or more ridges are specifically contemplated.

Figure 7 is a perspective view of a heat transfer element 14 in use within a blood vessel, showing only one helical lobe per segment for purposes of clarity. Beginning from the proximal end of the heat transfer element (not shown in Figure 7), as the blood moves forward during the systolic pulse, the first helical heat transfer segment 20 induces a counter-clockwise rotational inertia to the blood. As the blood reaches the second segment 22, the rotational direction of the inertia is reversed, causing turbulence within

the blood. Further, as the blood reaches the third segment 24, the rotational direction of the inertia is again reversed. The sudden changes in flow direction actively reorient and randomize the velocity vectors, thus ensuring turbulence throughout the bloodstream. During turbulent flow, the velocity vectors of the blood become more random and, in some cases, become perpendicular to the axis of the artery. In addition, as the velocity of the blood within the artery decreases and reverses direction during the cardiac cycle, additional turbulence is induced and turbulent motion is sustained throughout the duration of each pulse through the same mechanisms described above.

Thus, a large portion of the volume of warm blood in the vessel is actively brought in contact with the heat transfer element 14, where it can be cooled by direct contact rather than being cooled largely by conduction through adjacent laminar layers of blood. As noted above, the depth of the grooves 26, 30, 34 (Figure 4) is greater than the depth of the boundary layer which would develop if a straight-walled heat transfer element were introduced into the blood stream. In this way, free stream turbulence is induced. In the preferred embodiment, in order to create the desired level of turbulence in the entire blood stream during the whole cardiac cycle, the heat transfer element 14 creates a turbulence intensity greater than about 0.05. The turbulence intensity may be greater than 0.05, 0.06, 0.07 or up to 0.10 or 0.20 or greater.

Referring back to Figure 4, the heat transfer element 14 has been designed to address all of the design criteria discussed above. First, the heat transfer element 14 is flexible and is made of a highly conductive material. The flexibility is provided by a segmental distribution of bellows sections 25, 27 which provide an articulating mechanism. Bellows have a known convoluted design which provides flexibility. Second, the exterior surface area 37 has been increased through the use of helical ridges 28, 32, 36 and helical grooves 26, 30, 34. The ridges also allow the heat transfer element 14 to maintain a relatively atraumatic profile, thereby minimizing the possibility of damage to the vessel wall. Third, the heat transfer element 14 has been designed to promote turbulent kinetic energy both internally and externally. The modular or segmental design allows the direction of the invaginations to be reversed between segments. The alternating helical rotations create an alternating flow that results in mixing the blood in a manner analogous to the mixing action created by the rotor of a washing machine that switches directions back and forth. This mixing action is intended

to promote high level turbulent kinetic energy to enhance the heat transfer rate. The alternating helical design also causes beneficial mixing, or turbulent kinetic energy, of the working fluid flowing internally.

Figure 8 is a cut-away perspective view of an alternative embodiment of a heat transfer element 50. An external surface 52 of the heat transfer element 50 is covered with a series of axially staggered protrusions 54. The staggered nature of the outer protrusions 54 is readily seen with reference to Figure 9 which is a transverse cross-sectional view taken at a location denoted by the line 9-9 in Figure 8. In order to induce free stream turbulence, the height,  $d_p$ , of the staggered outer protrusions 54 is greater than the thickness of the boundary layer which would develop if a smooth heat transfer element had been introduced into the blood stream. As the blood flows along the external surface 52, it collides with one of the staggered protrusions 54 and a turbulent wake flow is created behind the protrusion. As the blood divides and swirls along side of the first staggered protrusion 54, its turbulent wake encounters another staggered protrusion 54 within its path preventing the re-lamination of the flow and creating yet more turbulence. In this way, the velocity vectors are randomized and turbulence is created not only in the boundary layer but throughout the free stream. As is the case with the preferred embodiment, this geometry also induces a turbulent effect on the internal coolant flow.

A working fluid is circulated up through an inner coaxial lumen 56 defined by an insulating coaxial tube 58 to a distal tip of the heat transfer element 50. The working fluid then traverses an outer coaxial lumen 60 in order to transfer heat to the exterior surface 52 of the heat transfer element 50. The inside surface of the heat transfer element 50 is similar to the exterior surface 52, in order to induce turbulent flow of the working fluid. The inner protrusions can be aligned with the outer protrusions 54, as shown in Figure 9, or they can be offset from the outer protrusions 54, as shown in Figure 8.

Figure 10 is a schematic representation of the invention being used to cool the brain of a patient. The selective organ hypothermia apparatus shown in Figure 10 includes a working fluid supply 10, preferably supplying a chilled liquid such as water, alcohol or a halogenated hydrocarbon, a supply catheter 12 and the heat transfer element 14. The supply catheter 12 has a coaxial construction. An inner coaxial lumen within the supply catheter 12 receives coolant from the working fluid supply 10. The coolant

travels the length of the supply catheter 12 to the heat transfer element 14 which serves as the cooling tip of the catheter. At the distal end of the heat transfer element 14, the coolant exits the insulated interior lumen and traverses the length of the heat transfer element 14 in order to decrease the temperature of the heat transfer element 14. The coolant then traverses an outer lumen of the supply catheter 12 so that it may be disposed of or recirculated. The supply catheter 12 is a flexible catheter having a diameter sufficiently small to allow its distal end to be inserted percutaneously into an accessible artery such as the femoral artery of a patient as shown in Figure 10. The supply catheter 12 is sufficiently long to allow the heat transfer element 14 at the distal end of the supply catheter 12 to be passed through the vascular system of the patient and placed in the internal carotid artery or other small artery. The method of inserting the catheter into the patient and routing the heat transfer element 14 into a selected artery is well known in the art.

Although the working fluid supply 10 is shown as an exemplary cooling device, other devices and working fluids may be used. For example, in order to provide cooling, freon, perflourocarbon, water, or saline may be used, as well as other such coolants.

The heat transfer element can absorb or provide over 75 Watts of heat to the blood stream and may absorb or provide as much as 100 Watts, 150 Watts, 170 Watts or more. For example, a heat transfer element with a diameter of 4 mm and a length of approximately 10 cm using ordinary saline solution chilled so that the surface temperature of the heat transfer element is approximately 5° C and pressurized at 2 atmospheres can absorb about 100 Watts of energy from the bloodstream. Smaller geometry heat transfer elements may be developed for use with smaller organs which provide 60 Watts, 50 Watts, 25 Watts or less of heat transfer.

The practice of the present invention is illustrated in the following non-limiting example.

#### Exemplary Procedure

1. The patient is initially assessed, resuscitated, and stabilized.
2. The procedure is carried out in an angiography suite or surgical suite equipped with fluoroscopy.
3. Because the catheter is placed into the common carotid artery, it is important to determine the presence of stenotic atheromatous lesions. A carotid duplex

- (doppler/ultrasound) scan can quickly and non-invasively make this determinations. The ideal location for placement of the catheter is in the left carotid so this may be scanned first. If disease is present, then the right carotid artery can be assessed. This test can be used to detect the presence of proximal common carotid lesions by observing the slope of the systolic upstroke and the shape of the pulsation. Although these lesions are rare, they could inhibit the placement of the catheter. Examination of the peak blood flow velocities in the internal carotid can determine the presence of internal carotid artery lesions. Although the catheter is placed proximally to such lesions, the catheter may exacerbate the compromised blood flow created by these lesions. Peak systolic velocities greater than 130 cm/sec and peak diastolic velocities > 100 cm/sec in the internal indicate the presence of at least 70% stenosis. Stenosis of 70% or more may warrant the placement of a stent to open up the internal artery diameter.
4. The ultrasound can also be used to determine the vessel diameter and the blood flow and the catheter with the appropriately sized heat transfer element could be selected.
  5. After assessment of the arteries, the patients inguinal region is sterilely prepped and infiltrated with lidocaine.
  6. The femoral artery is cannulated and a guide wire may be inserted to the desired carotid artery. Placement of the guide wire is confirmed with fluoroscopy.
  7. An angiographic catheter can be fed over the wire and contrast media injected into the artery to further to assess the anatomy of the carotid.
  8. Alternatively, the femoral artery is cannulated and a 10-12.5 french (f) introducer sheath is placed.
  9. A guide catheter is placed into the desired common carotid artery. If a guiding catheter is placed, it can be used to deliver contrast media directly to further assess carotid anatomy.
  10. A 10 f -12 f (3.3- 4.0 mm) (approximate) cooling catheter is subsequently filled with saline and all air bubbles are removed.
  11. The cooling catheter is placed into the carotid artery via the guiding catheter or over the guidewire. Placement is confirmed with fluoroscopy.
  12. Alternatively, the cooling catheter tip is shaped (angled or curved approximately 45 degrees), and the cooling catheter shaft has sufficient pushability and torqueability to be placed in the carotid without the aid of a guide wire or guide catheter.

13. The cooling catheter is connected to a pump circuit also filled with saline and free from air bubbles. The pump circuit has a heat exchange section that is immersed into a water bath and tubing that is connected to a peristaltic pump. The water bath is chilled to approximately 0° C.
- 5 14. Cooling is initiated by starting the pump mechanism. The saline within the cooling catheter is circulated at 5 cc/sec. The saline travels through the heat exchanger in the chilled water bath and is cooled to approximately 1° C.
15. It subsequently enters the cooling catheter where it is delivered to the heat transfer element. The saline is warmed to approximately 5-7° C as it travels along the inner  
10 lumen of the catheter shaft to the end of the heat transfer element.
16. The saline then flows back through the heat transfer element in contact with the inner metallic surface. The saline is further warmed in the heat transfer element to 12-15° C, and in the process, heat is absorbed from the blood, cooling the blood to 30° C to 32° C.
- 15 17. The chilled blood then goes on to chill the brain. It is estimated that 15-30 minutes will be required to cool the brain to 30 to 32° C.
18. The warmed saline travels back down the outer lumen of the catheter shaft and back to the chilled water bath where it is cooled to 1° C.
19. The pressure drops along the length of the circuit are estimated to be 2-3  
20 atmospheres.
20. The cooling can be adjusted by increasing or decreasing the flow rate of the saline. Monitoring of the temperature drop of the saline along the heat transfer element will allow the flow to be adjusted to maintain the desired cooling effect.
21. The catheter is left in place to provide cooling for 12 to 24 hours.
- 25 22. If desired, warm saline can be circulated to promote warming of the brain at the end of the therapeutic cooling period.

While the particular invention as herein shown and disclosed in detail is fully capable of obtaining the objects and providing the advantages hereinbefore stated, it is to  
30 be understood that this disclosure is merely illustrative of the presently preferred embodiments of the invention and that no limitations are intended other than as described in the appended claims.



## CLAIMS

We claim:

1           1.     A selective organ heat transfer device, comprising:  
2           a flexible catheter capable of insertion to a selected feeding artery in the vascular  
3           system of a patient;  
4           a heat transfer element attached to a distal end of said catheter;  
5           a plurality of exterior surface irregularities on said heat transfer element, said  
6           surface irregularities being shaped and arranged to create turbulence in  
7           surrounding fluid, said surface irregularities having a depth at least equal  
8           to the boundary layer thickness of flow in the feeding artery; and  
9           an inner coaxial tube disposed within said heat transfer element, said inner  
10          coaxial tube being connected in fluid flow communication with an inner  
11          coaxial tube within said catheter.

1           2.     A heat transfer device as recited in claim 1, wherein said heat transfer  
2          element comprises a bellows.

1           3.     A heat transfer device as claimed in claim 1, wherein said heat transfer  
2          element comprises a plurality of heat transfer segments, and further comprising a flexible  
3          joint connecting each of said heat transfer segments to adjacent said heat transfer  
4          segments.

1           4.     A heat transfer device as recited in claim 3, wherein said flexible joint  
2          comprises a bellows.

1           5.     A heat transfer device as recited in claim 3, wherein:  
2           said surface irregularities comprise a helical ridge and a helical groove formed on  
3           each said heat transfer segment; and  
4           said helical ridge on each said heat transfer segment has an opposite helical twist  
5           to said helical ridges on adjacent said heat transfer segments.

1           6.     A heat transfer device as recited in claim 1, wherein:  
2           said surface irregularities comprise protrusions on said exterior surface; and  
3           said protrusions are axially staggered along said exterior surface.

1           7.     A heat transfer device as recited in claim 1, further comprising a plurality  
2           of interior surface irregularities in said heat transfer element, said interior surface  
3           irregularities being shaped and arranged to create turbulence in fluid within said heat  
4           transfer element, said interior surface irregularities having a depth at least equal to the  
5           boundary layer thickness of flow within said heat transfer element.

1           8.     A heat transfer device as recited in claim 7, wherein:  
2           said heat transfer element comprises a plurality of heat transfer segments;  
3           said interior surface irregularities comprise a helical ridge and a helical groove  
4           formed within each said heat transfer segment; and  
5           said helical ridge within each said heat transfer segment has an opposite helical  
6           twist to said helical ridges within adjacent said heat transfer segments.

1           9.     A heat transfer device as recited in claim 7, wherein:  
2           said surface irregularities comprise protrusions on said interior surface; and  
3           said protrusions are axially staggered along said interior surface.

1           10.    A selective organ heat transfer device comprising:  
2           a flexible coaxial catheter capable of insertion to a selected feeding artery in the  
3           vascular system of a patient;  
4           an articulated heat transfer element attached to a distal end of said catheter, said  
5           heat transfer element having an exterior diameter substantially less than  
6           the inner diameter of the selected feeding artery;  
7           a plurality of heat transfer segments on said heat transfer element, each said  
8           segment having a plurality of exterior surface irregularities, said surface  
9           irregularities being shaped and arranged to create turbulence in  
10          surrounding fluid, said surface irregularities having a depth at least equal  
11          to the boundary layer thickness of flow in the feeding artery;  
12          a flexible joint connecting each of said heat transfer segments to adjacent said  
13          heat transfer segments; and  
14          an inner coaxial tube disposed within said heat transfer element, said inner  
15          coaxial tube being connected in fluid flow communication with an inner  
16          coaxial tube within said catheter.

1           11.    A heat transfer device as recited in claim 10, wherein said flexible joint  
2    comprises a bellows.

1           12.    A heat transfer device as recited in claim 10, wherein:  
2           said surface irregularities comprise a helical ridge and a helical groove formed on  
3           each said heat transfer segment; and  
4           said helical ridge on each said heat transfer segment has an opposite helical twist  
5           to said helical ridges on adjacent said heat transfer segments.

1           13.    A method for selectively controlling the temperature of a selected organ of  
2 a patient, said method comprising:  
3           providing a catheter having a heat transfer element attached to a distal end  
4           thereof, said heat transfer element having a plurality of exterior surface  
5           irregularities, said surface irregularities having a depth greater than the  
6           boundary layer thickness of flow in the feeding artery of the selected  
7           organ;  
8           inserting said catheter through the vascular system of the patient to place said heat  
9           transfer element in the feeding artery of the selected organ;  
10          creating turbulence around said surface irregularities at a distance from said heat  
11          transfer element greater than the boundary layer thickness of flow in the  
12          feeding artery, thereby creating turbulence throughout the blood flow in  
13          the feeding artery;  
14          circulating fluid into said heat transfer element via an internal lumen of said  
15          catheter and via an internal lumen of said heat transfer element;  
16          circulating fluid out of said heat transfer element via an external lumen of said  
17          heat transfer element; and  
18          transferring heat between said heat transfer element and the blood in the feeding  
19          artery to selectively control the temperature of the selected organ.

1           14.    A method as recited in claim 13, wherein:  
2           said surface irregularities on said heat transfer element comprise a plurality of  
3           segments of helical ridges and grooves having alternating directions of  
4           helical rotation; and  
5           turbulence is created by establishing repetitively alternating directions of helical  
6           blood flow with said alternating helical rotations of said ridges and  
7           grooves.

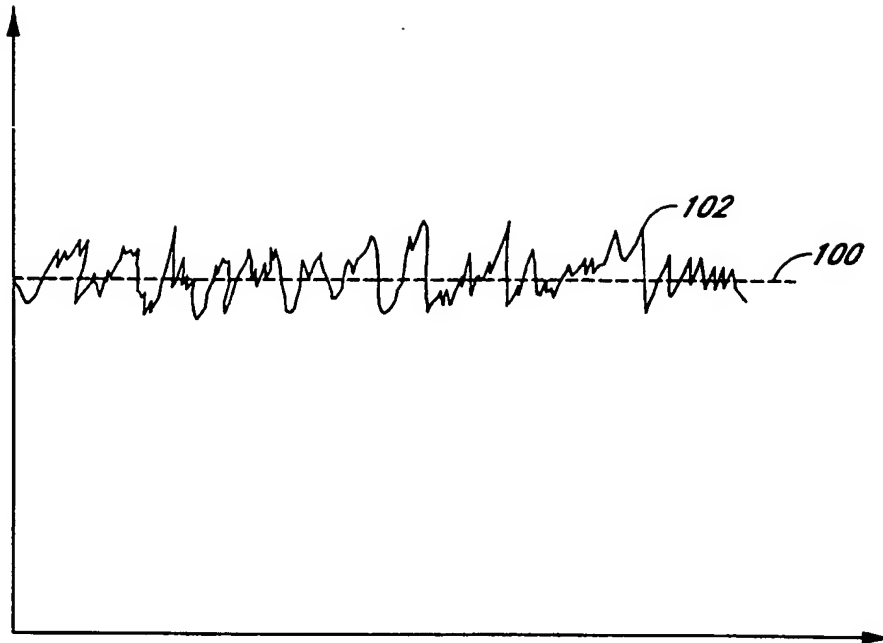


Fig. 1

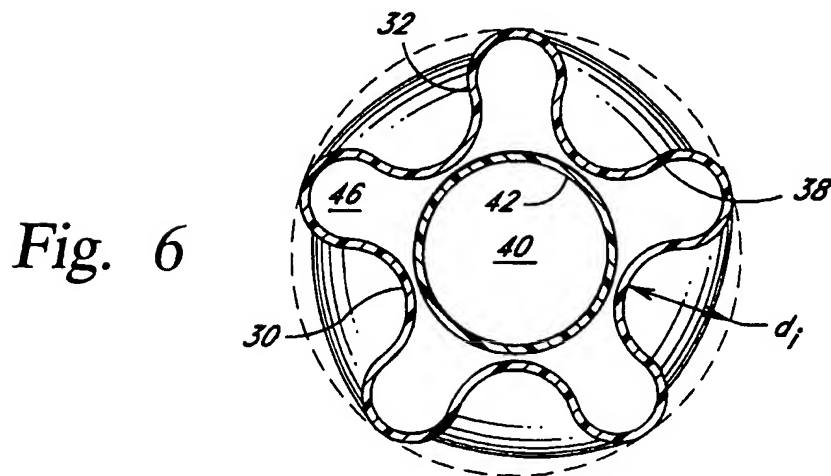


Fig. 6

2/6

Fig. 2A

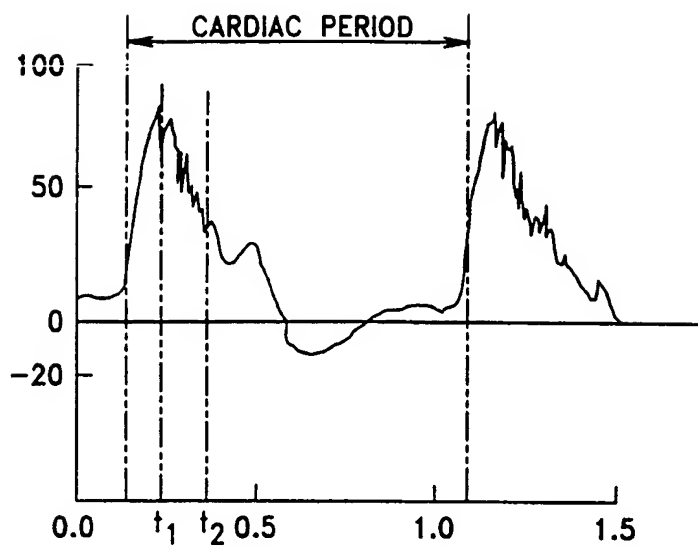


Fig. 2B

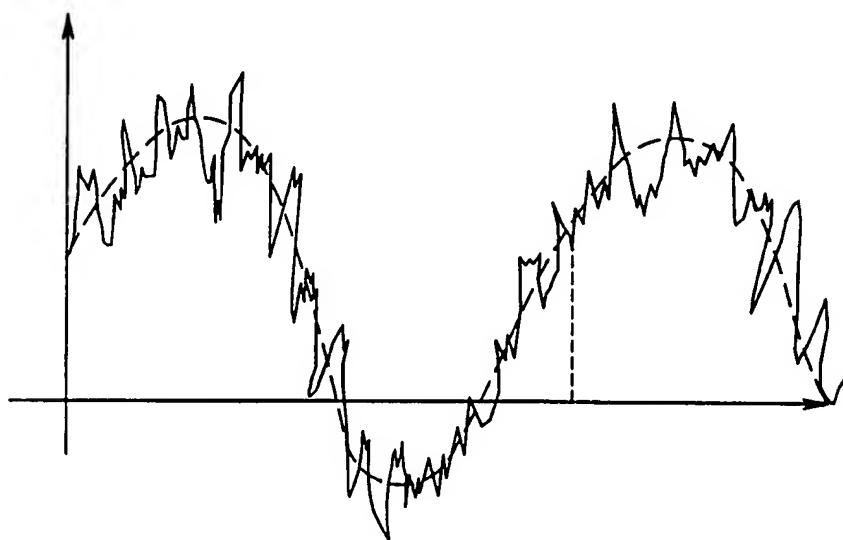
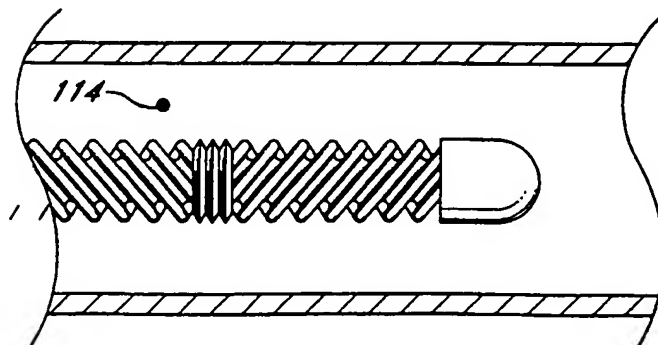


Fig. 2C



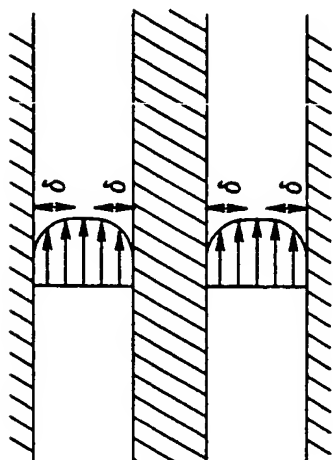


Fig. 3C

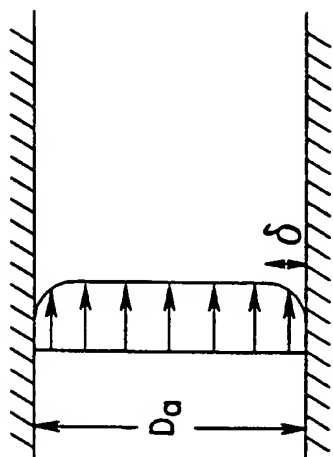


Fig. 3B

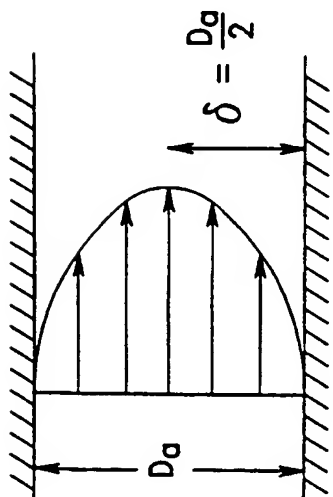


Fig. 3A

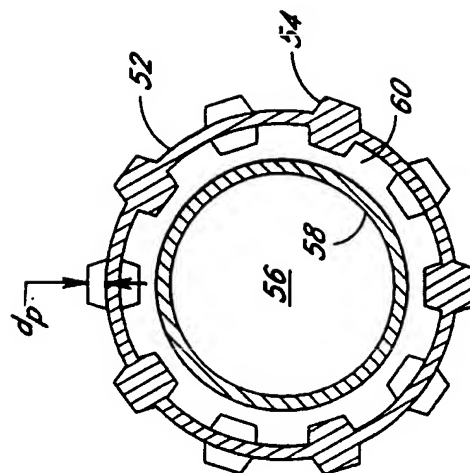


Fig. 9

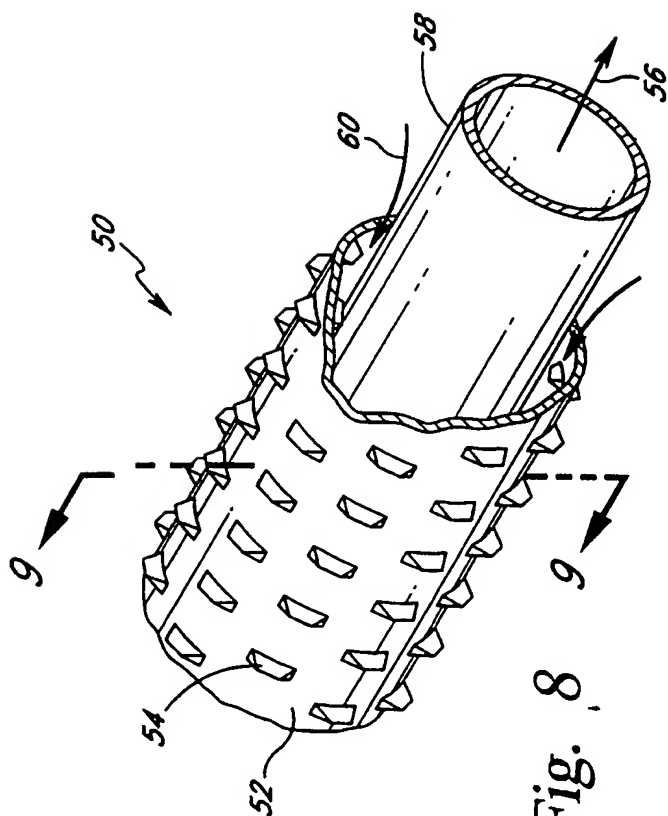


Fig. 8

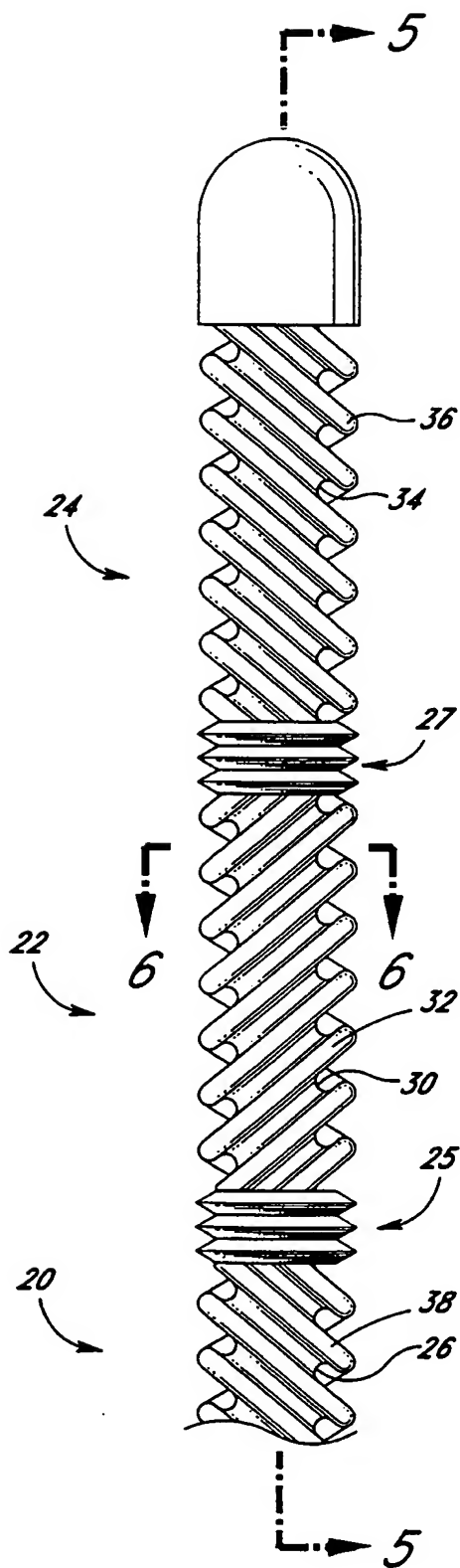


Fig. 4

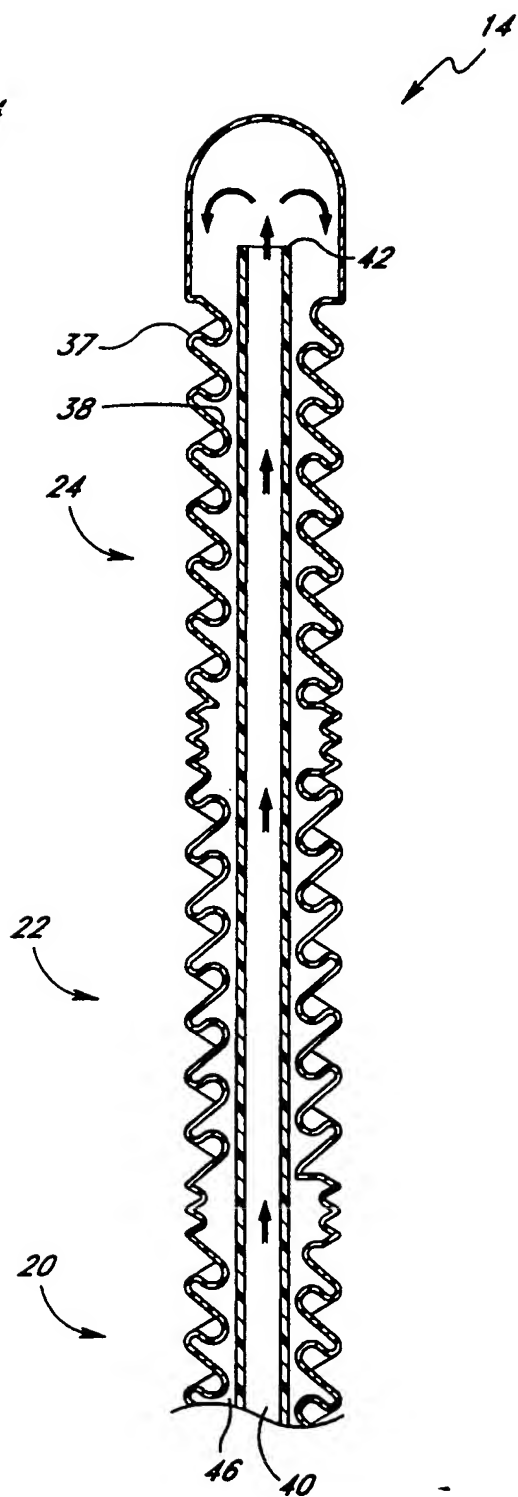
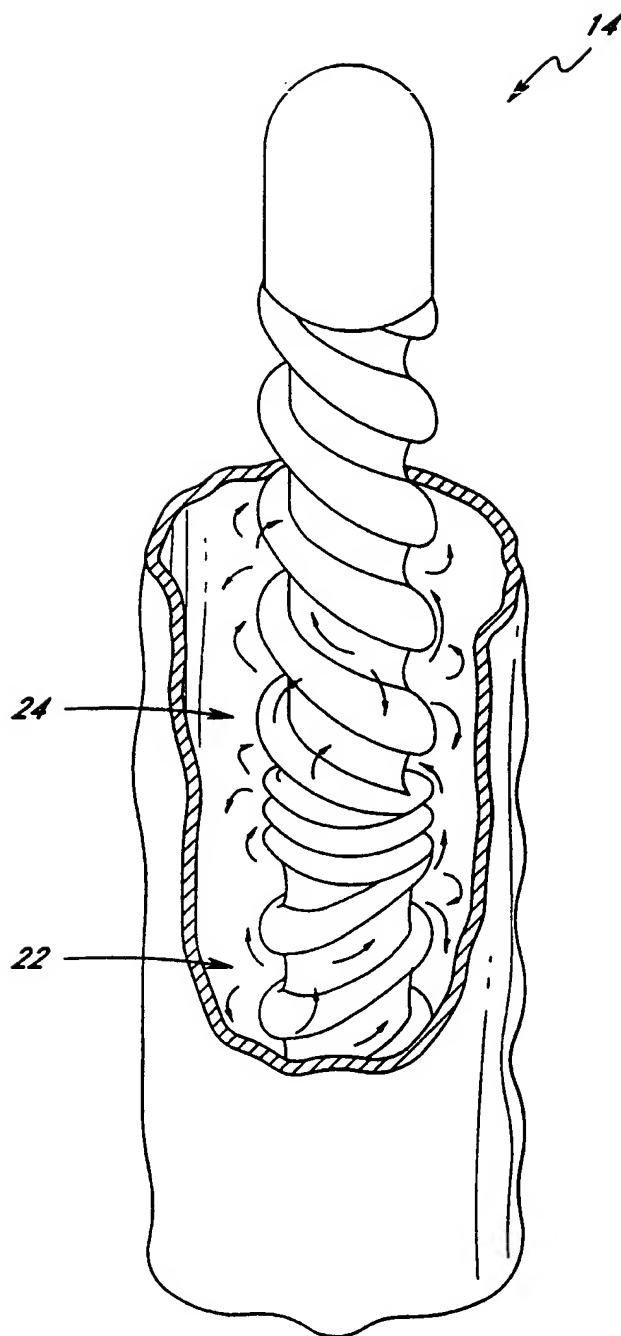
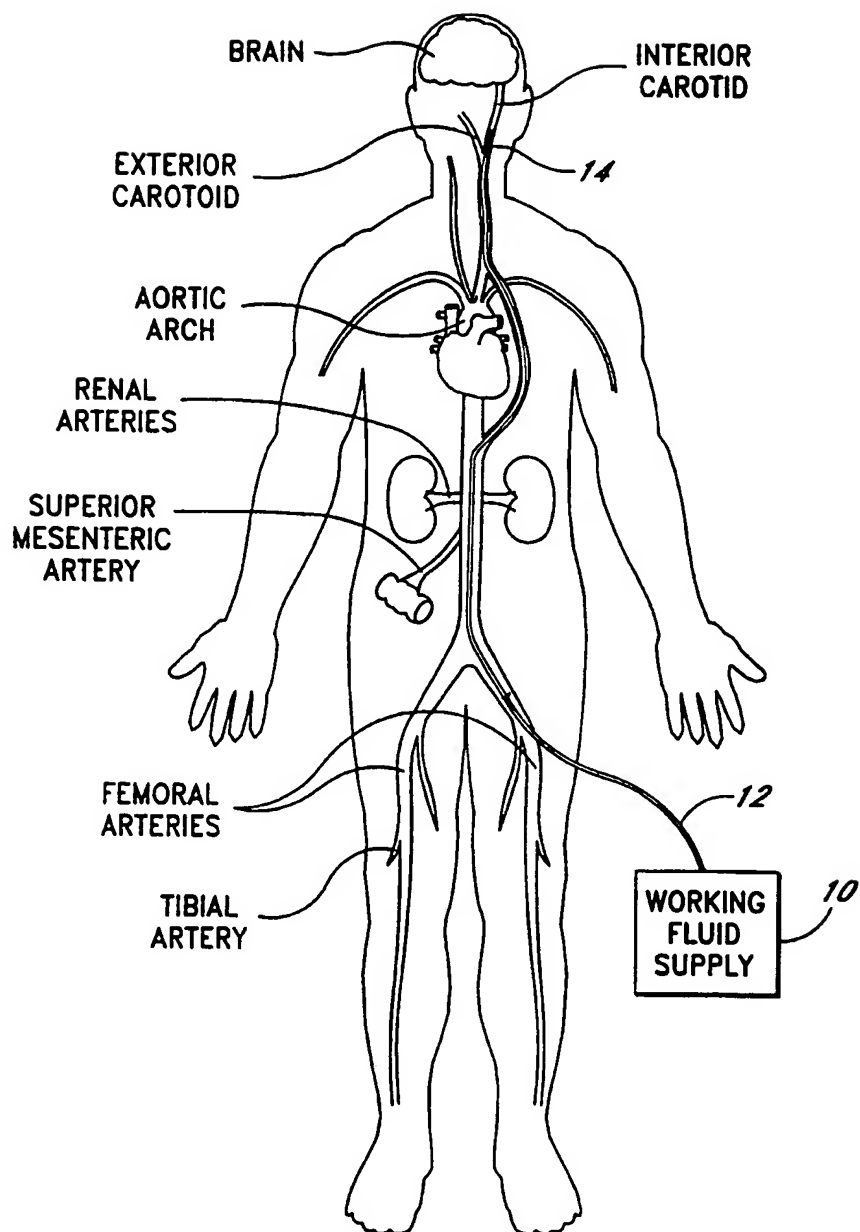


Fig. 5



*Fig. 7*

*Fig. 10*

## INTERNATIONAL SEARCH REPORT

International application No.  
PCT/US99/06265

## A. CLASSIFICATION OF SUBJECT MATTER

IPC(6) :A61F 7/00

US CL :607/104

According to International Patent Classification (IPC) or to both national classification and IPC

## B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)

U.S. : 604/239, 264, 916; 607/104-106

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practicable, search terms used)

## C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category*	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
X	US 4,190,033 A (FOTI) 26 February 1980, Figs. 8-10.	1-4, 10-12
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Y		6-9, 13, 14
Y	US 5,624,392 A (SAAB) 29 April 1997, Fig. 6.	5

☐ Further documents are listed in the continuation of Box C. ☐ See patent family annex.

* Special categories of cited documents:	*T* later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention
*A* document defining the general state of the art which is not considered to be of particular relevance	*X* document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone
*E* earlier document published on or after the international filing date	*Y* document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art
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Name and mailing address of the ISA/US  
Commissioner of Patents and Trademarks  
Box PCT  
Washington, D.C. 20231

Facsimile No. (703) 305-3230

Authorized officer

ROSALIND KEARNEY

Telephone No. (703) 308-2711